

10/522535

WO 2004/013276

Josef PRASSLER et al.
"Novel Tricistronic Vectors and Uses Therefor"
Attorney Docket No. 37629-0064US

PCT/IB2003/003681

1/25

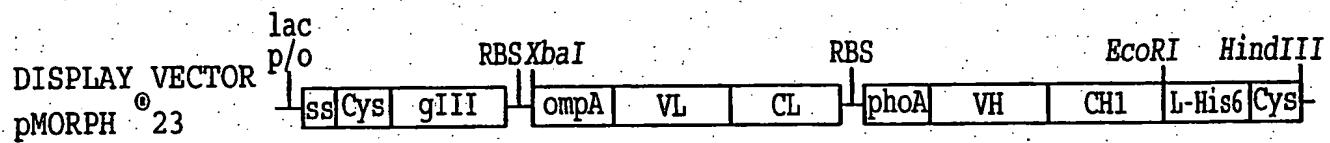


FIG. 1

2/25

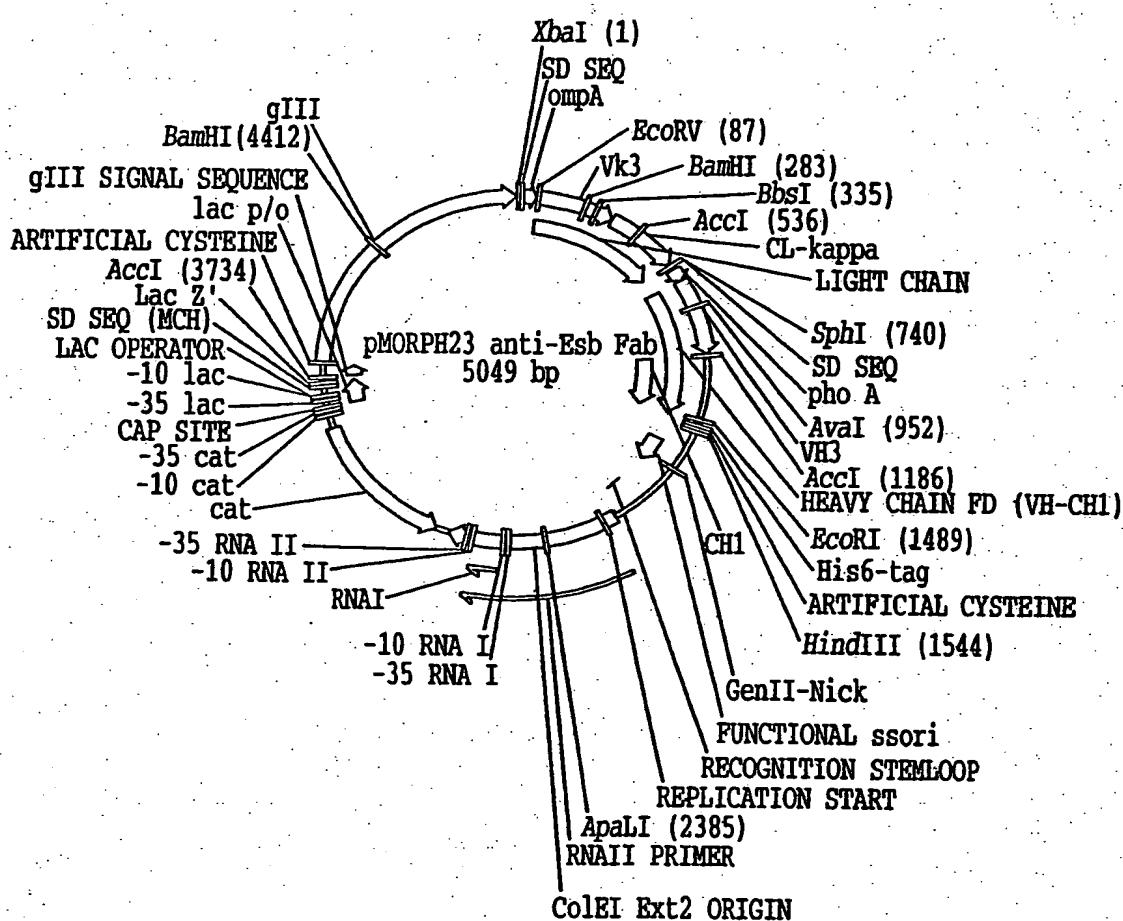


FIG. 2A

10/522535

WO 2004/013276

Josef PRASSLER et al.
 "Novel Tricistronic Vectors and Uses Therefor"
 Attorney Docket No. 37629-0064US

PCT/IB2003/003681

3/25

BamI

EcoRI

EcoRV

1 CTAGATAACG AGGGCAAAA ATGAAAAAGA CAGCTATCGC GATTGCAGTG GCACTGGCTG GTTTCGCTAC CGTAGCCGAG GCGGATATCG TGCTGACCCA
 101 GATCTATTGC TCCCGTTTT TACTTTTCT GTCCGATAGCG CTAACGTAC CGTGACCGAC CAAAGCGATG GCATCGCGTC CGCTATAGC ACGACTGGT
 101 GAGCCGGCG ACCCTGAGCC TGTCTCCGGG CGAACGTGCG ACCCTGAGCT GCAGAGCGAG CCAGCTGTG TTCTCGTTCT ATCTGGCTTG GTACCAGCAG
 201 CTGGGCCGC TGGAACCTGG ACAGAGGCC CGCTTGACCG TGGAACCTGA CGTCTCGCTC GGTCAGACAA AGAGCAAGAA TAGACCCGAAAC CATGGTGTGTC
 201 AAACCAAGTC AAGCACCGCG TCTATTAATT TATGGTGTCTT CTGTCGTCG AACTGGGGTC CGGCGCGTT TTAGGGCTC TGGATCCGGC ACGGATTAA
 TTTGGTCCAG TTCGTGGCGC AGATAATTAA ATACCAACGAA GAGCAGCAAG TTGACCCCGAG GGCGCGCAA AATCGCCGAG ACCTAGGCCG TGCTAAAT
 BbsI MscI

301 CCCTGACCAT TAGCAGCCTG GAACCTGAAG ACTTTCGAC TTATTATTC CAGCAGCGTG GTAATTATTC TATTACCTT GGCCAGGGTA CGAAAGTGA
 GGGACTGGTA ATCGTCGAC CTGGACTTC TGAAACGCTG AATAATAACG GTGTCGAC CATTATAAG ATAATGGAAA CGGTCCCCAT GCTTCAACT
 BsiNI

401 AATTAACGT ACGGTGGCTG CTCCGAGCGT GTTTATTTT CCGCCGAGCG ATGAAACAAT GAAAAGCGGC ACGGCGAGGG TGGTGTGCT GCTGAAACAC
 TTAATTGCA TGCCACCGAC GAGGCTCGCA CAAATAAAA GCGCGCTCGC TACTTGTGA CTTTTCGCG TGCGCTCGC ACCACACGGG CGACTTGTG
 501 TTTTATCCGC GTGAAGCGAA AGTTCAGTGG AAAGTAGACA ACGGCGCTGCA AAGCGGCAAC AGCCAGGAAA GCGTGCACCGA ACAGGATAGC AAAGATAGCA
 AAAATAGCG CACTTCGTT TCAAGTCACC TTTCATCTGT TGCGCGCTG TTGGTCTTT CGCACTGGCT TGTCTATCG TTCTATCGT
 601 CCTATTCTCT GAGCAGCACC CTGACCCCTGA GCAAGCGGA TTATGAAAAA CATAAAAGTGT ATGGTGGGA AGTGAACCAT CAAGGTCTGA GCAGCCGGT
 GGATAAGAGA CTCGTGTGG GACTGGGACT CGTTTCGCTT AATACTTTT GTATTTCACA TACGCACCGT TCACTGGTA GTTCCAGACT CGTCGGGCCA

EcoRI for 100.0%

StuI

SphI

SapI

701 GACTAAATCT TTTAATCGTG GCGAGGGCTG ATAAGCATGC GTAGGAGAAA ATAAATGAA ACAAAAGCACT ATTGCACTGG CACTCTTACG GTTGCCTTC
 CTGATTTAGA AAATTAGCAC CGCTCCGGAC TATTCGTAGC CATCCTCTT TATTTACTT TGTTTCGIGA TAACGTGACCG GTGAGAATGG CAACGAGAAG
 EcoRI for 100.0% MfeI

801 ACCCTCTTA CCAAAGGCCA GGTGCAATTG GTGAAAGCG GGGCGCCCT GGTGCAACCG GGGGGCAGCC TCGGTCTGAG CTGGCCGGCC TCCTGGATTA
 TGGGACAAT GTTTGGGT CCACGTTAAC CACCTTCGCG CGCGCGCGA CCACGTTGGC CGCGCGTGG AGCGAGACTC GACGCGCGGG AGGCCTAAAT
 901 CCTTTCTTC TTATGGTGGT AAATGGTGC GCAAGCCCC TGGGAAGGGT CTGGAGTGG TGAGGGTAT CCATTATTCT GGTAGCTCTA CCTTATTATGC
 GAAAAGAAG AATACCACCA TTAACCCACG CGGTTGGGG ACCCTTCCCA GAGCTCACCC ACTGCCATA GGTAAATAAGA CCATCGAGAT GGATAATACG
 1001 GGATAGCGTG AAAGCCGTT TTACCATTT ACCTGATAAT TCGAAAACA CCTCTATCT GCAATGAAAC AGCTGGCTG CGGAAGATAC GGGCGTGTAT
 CCTATCGCAC TTCCGGCAA AATGGTAAAG TGCACCTATA AGCTTTTGT GGGACATAGA CGTTTACTTG TCGGACCCAC GCCTCTATG CGCCACACATA

SalI

BssHII

SapI

StyI

BpuI

1101 TATTGCGCGC GTGCTCTTCA TAATGCGGT GTTGTGGGTT TTGATCATTG GGGCAAGGC ACCCTGGTG CGGGTAGCTC AGGGTGCAGCC AAAGGTCAA
 ATAACCGCGC CACGAGAACT ATTACCCCGA CCAACCCAA AACTAGTAAC CCCGGTCCG TGGGACACT GCGAATCGAG TCGCAGCTGG TTTCAGGTT
 1201 GCGTGTCTCC GCTGCGCTCC AGCAGCAAAA GCAACCGGG CGGCACGGCT GCGCTGGGT GCGTGGTAA AGATTATTTC CGGAACCCAG TCACCGTGG
 CGCACAAAG CGACCGAGGC TCGTGTCTT CGTGGTCCG GCGGTGCCGA CGGGACCGA CGGACCAATT TCTAATAAG GGCGCTGGTC AGTGGCACTC
 1301 CTGGACACGC GGGCGCTGA CCACCGCGT GCATACCTT CGCGCGCTGC TGCAAAAGCAG CGGCGCTGTAT AGCCTGAGCA CGGTGTGAC CGTGGCGAGC
 GACCTGTGCG CCCCGCGACT GGTGCGCGA CGTATGGAAA GGGCGCCACG AGCTTCGTC GCGGACATA TCGGACTCGT CGCAACACTG CGACCGCTGG

EcoRI

1401 AGCAGCTTAG GCACTCAGAC CTATATTTC AACGTGAACC ATAAACCGAG CAACACCAA GTGGATAAAA AAGTGGAAAC GAAAAGCGAA TTCCCAGGG
 TCGTCAATC CGTGAAGCTG GATATAAAAG TTGCACTTGG TATTTGGCTC GTGTTGGTT CACCTATTTC TTGACCTTGG CTTTTGGCTT AAGGGTCCCC
 BssHII

Asci

KpnI

4/25

INCL rev 100.0

1501 GGAGCGGAGG CGCGCCGCAC CATCATCACC ATCACTGCTG ATAACCTTGA CCTGTGAAGT GAAAAATGGC GCAGATTTG CGACATTIT TTGTCGCC
 CCTCGCCCTGC CGCGCGCGTG GTAGTAGTGG TAGTGACGAC TATTGAACT GGACACTTCA CTTTTAACCG CGCTTAACAC GCTGAAAGA AAACAGAOGG
 1601 GTTTAATGAA ATTGTAACG TTAATAATT TTGTAACATT GCGTAAATT TTTGTTAAAT CAGCTATT TTAAACCAAT AGGCGAAT CGCAAATC
 CAAATTACTT TAACATTGCA AATTATAAAA CAATTTAAG CGCAATTAA AAACAATTAA GTGAGTAA AAATGGTAA TCCGGCTTAA GCGTTTACG
 1701 CCTTATAAAT CAAAAGAATA GACCGAGATA GGTTGAGTG TGTTGCACTG TTGGAACAAG AGTCCACTAT TAAAGAACGT GGACTCCAC GTCAAAGGGC
 GGAATATTAA GTTTCTTAT CTGGCTCTAT CCCAACTCAC ACAAGGTCA AACCTTGTTC TCAGGTATA ATTCTTGA CCGTGGGTG CAGTTCCCG
 1801 GAAAAACCGT CTATCAGGGC GATGGCCCAC TACCGAGAAC ATCACCTAA TCAAGTTTT TGGGGTCAG GTGCGTAA GCACAAATC GGAACCCCTAA
 CTTTGGCA GATAGTCCCG CTACGGGTG ATGCTTGG TAGTGGATT AGTCAAAA ACCCCAGCTC CACGGCATT CGTGAATTAG CCTTGGGATT
 1901 AGGGAGCCCC CGATTTAGAG CTGACGGGG AAAGCGGCG AACCTGGCA GAAAGGAAGG AAAGGAGGG GCGCTAGGGC GCTGGCAAGT
 TCCCTGGGG GCTAAATCTC GAACIGCCCC TTTCGGCCGC TTGACCCGCT CTTCTTGC CTTCTCGCC CGCGATCCCG CGACCGTCA

NheI

2001 GTAGCGGTCA CGCTCGCGCT AACACCACAA CCCGCGCGGC TTAATGGGCC GCTACAGGG CGCTGCTAGC CAATGAGCA AAAGGCCAG AAAAGGCCAG
 CATCGCCAGT CGGAGCGCA TTGGTGGTGT GGCGCGCGC AATTACGCGG CGATGTCGG CGACACTCGT TTTCGGTGTG TTTCGGTC
 2101 GAACCGTAAA AAGGCCGCGT TGCTGGGTGTT TTCCATAGG CTCCGCCCCC CTGACGAGCA TCACAAAAT CGACGCTCAA GTCAAGGGTGC GCGAAACCCG
 CTTGGCATTT TTCCGGCGCA ACAGACCGAA AAAGGTATCC GAGGGGGGG GACTGCTGT AGTGTGTTTA GCTGCGAGT CAGTCTCAC CGCTTGGC
 2201 ACAGGACTAT AAAAGATACCA GGCGTTTCCC CCTGGAAGCT CCTCTGTCGG CTCTCTGTT CGGACCTGG CGCTTACCGG ATACCTGTC GCCTTCTC
 TGTCCTGATA TTCTATGGT CGCGAAAGGG GGACCTTCGA GGGAGCACG GAGAGGACAA GGCTGGGAC GCGAATGGCC TATGGACAGG CGGAAAGAGG
 2301 CTTCGGGAAG CGTGGCGCTT TCTCATAGCT CACCGCTGTAG GTATCTCAGT CGGCTGTAGG TGGTGTAGG TGGTGTAGC AACCCCCGT
 GAAGCCCTTC GCACCGCGA AGAGTATCGA GTGCGACATC CATAGAGTCA AGCCACATCC AGCAAGOGA GTGCGACCCG ACACACGTG TGCGGGGCA
 2401 TCAGCTGGAC CGCTGGCCT TATCGGTAA CTATGCTCTT GAGTCAACCC CGGTAAGACCA CGACTTATCG CCACTGGCAG CAGCACCTGG TAACAGGATT
 AGTCAGGCTG GCGAGCGGA ATAGGCCATT GATAGCAGAA CTCAAGGTTGG GCCATTCTGT GCTGAATAGC GTGACCGTGTG ATGCTCTAA
 2501 AGCAGAGCGA GGTATGTAGG CGGTGCTACA GAGTTCTGAA AGTGGTGGCC TAACATCGC TACACTAGAA GAACAGTATT TGGTATCTGC GCTCTGCTGT
 TCGTCTGCT CCATACATCC GCGACCGATGT CTCAAGAACT TCACCAACCG ATTGATGCCG ATGATGCTT CTGTCATAA ACCATAGACG CGAGACGACA
 2601 AGCCAGTIACT TTTCGGAAA AGAGTTGGTA GCTCTGATC CGGCAAACAA ACCACCGCTG GTAGGGTGG TTTTTTGTG TGAAGGAGC AGATTACGG
 TCGGTCAATG GAAGCTTTT TCTCAACCCT CGAGAACATG GCGTTTGTG TGTGGCGAC CATGCCACC AAAAACACAA ACCTGTCG TCTAATGCC

BglII

2701 CAGAAAAAA GGATCTCAAG AGATCTTT GATCTTTCTT ACAGGGTCTG ACGCTCAGTG GAAACAAAAC TCAAGTTAAG GGATTTGGT CAGATCTAGC
 GTCTTTTTT CCTAGAGTTC TTCTAGAAA CTAGAAAAGA TGCCCCAGAC TGCGAGTCAC CTTGCTTTTG AGTCAATTTC CCTAAACCA GTCTAGATCG
 2801 ACCAGCGCTT TAAGGGCACC AATAACTGCC TTAAAAAAAT TACGCCCGC CTCGCACTC ATCGCAGTAC TGTGTAATT CATTAAAGCAT TCTGCCGACA
 TGGTCCGCAA ATTCCCGTGG TTATTGACGG AATTTTTTTA ATGGGGGGG GGAAGGTGAG TAGGTCATG ACAACATTAA GTAATTGCTA AGACGGCTGT
 2901 TGGAAGCCAT CACAAACCGC ATGATGAAAC TGAATCGCA GGGCATCAG CACCTTGCG CTTGCGTAT AATATTGCC CATACTGAAA AGCGGGCGA
 ACCTTCGGTA GTGTTTGCCTG TACTACTTGG ACTTAGCGGT CGCGCTAGTC GTGGAACAGC GGAACGCATA TTATAAACCG GTATCACTT TGGCCCGCT
 3001 AGAAGTGTCA CATATGGT ACGTTAAAT CAAAACGGT GAAACTCACC CAGGGATTGG CTGAGACGAA AAACATATT CAAATAAACCTT TAGGAA
 TCTTCACACG GTATAACCGA TGCAAATTAA GTTTGACCA CTTTGAGTGG GTCCCTAAC GACTCTGCTT TTGTTATAAG AGTATTGAA GAAATCCCT
 3101 ATAGGCCAGG TTTCACCGT AACACGCCAC ATCTTGCGAA TATATGTGTA GAAACTGCCG GAAATCGTC TGGTATTCACTC CTCAGAGGGA TGAAGACGGT
 TATCCGGTCC AAAAGTGGC TTGTCGGTG TAGAACGCTT ATATACACAT CTTTGACGC CTTAGCAGC ACCATAAGTG AGGCTCGCT ACTTTTGC
 3201 TCAGTTGCT CATGAAACG GGTGTAACAA GGGTGAACAC TATCCCATAT CACCAAGCTCA CGCTCTTCA TGGCCATAGG GAACCTGGG TGAGCATICA
 AGTCAAACGA GTACCTTTG CCACATGTT CCCACTTGTG ATAGGGTATA GTGGTCGAGT GGCAGAAAGT AACGGTATGC CTTGAGGCC ACTCGTAAGT
 3301 TCAGGGGGC AAGAATGTA ATAAAGGCCG GATAAAACTT GTGCTTATT TTCTTACGG TCTTAAAGA GGCCTAATA TCCAGCTGAA CGCTCGGTT
 AGTCCGCCCG TTCTTACACT TATTTCGGC CTATTTGAA CACGAATAAA AAGAAATGCC AGAAATTTC CCGGCAATT AGGTGACTT GCGAGACCAA
 3401 ATAGGTACAT TGAGCAACTG ACTGAAATGC CTCAAATGT TCTTACGAT GCCATTGGGA TATACAAAC GGGTATATC CAGTGAATT TTCTCATT
 TATCCATGTA ACTCGTGTGAC TGACTTACG GAGTTTACA AGAAATGCTA CGGTAACCC ATAGTGTG CACCATATAG GTCACTAAAA AAAGAGGAA

AatII

3501 TTAGCTTCT TAGCTCTGAA AATCTCGAT AACTCAAAA ATACGCCCG TAGTGATCTT ATTCATTAT GTGAAAGT GGAACCTCAC CGACCGCTA
 AATCGAAGGA ATCGAGGACT TTAGAGCTA TTGAGTTTT TATGCGGGC ATCACTAGAA TAAAGTAATA CCACTTCAA CCTTGGAGTG GCGTGCAGAT

M13 rev 100.0

FIG. 2B-2

5/25

3601 ATGTGAGTTA GCTCACTCAT TAGGCACCCC AGGCTTACA CTTATGCTT CGCGCTGTA TGTTGTTGAA AATTGAGC GGATAACAAT TTACACAGG
 TACACTCAAT CGAGTGAGTA ATCCGTGGG TCCGAAATGT GAATACGAA GGCAGGACAT ACAACACACC TTAACACTCG CCTATTGTTA AAGTGTGTC
 M13 rev 100.0

3701 AAACAGCTAT GACCATGATT ACGAAATTCT AGTATACGAG GGCACAAAAT GAAAAAACTG CTGTTGCGA TTCCGCTGGT GTGCCGTC TATAGCCATA
 TTGTCGATA CTGGTACTAA TGCTTAAAGA TCATATGTC CCGTTTTTA CTTTTTGAC GACAAGCGT AAGGGACCA CCACGGCAAG ATATCGGTAT
 3801 GCGACTACTG CGACATCGAG TTGCGAGAA CAGTTGAAAG TGTTTAGCA AAACCCATA CAGAAAATTG ATTACTAAC GTCCTGAAAG ACGACAAAC
 CGCTGATGAC GCTGTAGTC AAACGCTTT GTCAACTTTC ACAAAATCGT TTGCGGTAT GTCTTTAAG TAAATGATG CAGACCTTC TGCTGTTTG
 3901 TTGAGATGTT TACGCTAACT ATGAGGGCTG TCTGTTGAAT GTCAACAGGCG TTGAGTTTG TACTGGTGAC GAAACTCAGT GTTACGGTAC ATGGGTTCT
 AAATCTAGCA ATGCGATGTA TACTCCGAC AGACACCTTA CGATGTCGC AACATCAAAC ATGACCACTG CTTTGAGTC CAATGCCATG TACCCAGGA
 4001 ATGGGCTTG CTATCCCTGA AAATGAGGGT GGTGGCTCTG AGGGTGGCGG TTCTGAGGGT GGCGGCTCTG AGGGTGGCGG TACTAAACCT CCTGAGTACG
 TAACCGAAC GATAAGGACT TTGACTCCCA CCACCGAGAC TCCCACCGC AAGACTCCA CGCCCGAGAC TCCCACCGCC ATGATTGGA GGACTCATGC
 4101 GTGATACACC TATCCGGC TATACTTATA TCAACCCCTCG ACAGGGCACT TATCCGCTG GTACTGAGCA AAACCCCGT AATCCTAAC TCTCTCTIGA
 CACTATGTTG ATAAGGGCGG ATATGAATAT AGTTGGGAGA GCTGCGTGA ATAGGGCGAC CATGACTCGT TTGGGGCGA TTAGGATTAG GAAGAGAACT
 4201 GGAGCTCTAG CCTCTTAATA CTTCATGTT TCAGAATAAT AGGTTCGAA ATAGGCAGGG GGCAATTAACT GTTATACGG GCAGTGTAC TCAAGGCACT
 CCTCAGACTC GGAGAAATTGAAAGTACAA AGTCTTATTA TCCAAGGCTT TATCCGCTCC CGTAATTGA CAAATATGCC CGTGACAAATG AGTCCGTCGA
 4301 GACCCGTTA AAACCTTATTA CGAGTACACT CCTGTTATCAT CAAAGGCAT GTATGAGCT TACTGGAAACG GTAAATTGAG AGACTGGCT TTCCATTCTG
 CTGGGCAAT TTGATAAT GGTGATGTA GGACATAGTA GTTTGCGTA CATACTGCGA ATGACCTTG CATTAACTC TCTGACCCGA AAGGTAAGAC
 4401 GCTTAATGA GGATCCATTG TTGTTGAAAT ATCAAGGCCA ATGCTCTGAC CTGCTCAAC CTCTGTCAA TGCTGGCGG GGCTCTGGTG GTGGGTCGG
 CGAAATTACT CCTAGGTAAG CAAACACTTA TAGTTCCGGT TAGCAGACTG GACGGAGTT GAGGACAGTT ACACGGCGG CGAGACCCAC CACCAAGAAC
 4501 TGGCGCTCT GAGGGTGGCG GCTCTGAGGG TGCGGTTCT GAGGGTGGCG GCTCTGAGGG TGCGGTTCC GGTGGGGCT CGGGTTCCGG TGATTTGAT
 ACCGCCGAGA CTCCACCGC CGAGACTCCC ACCGCCAAGA CTCCACCGC CGAGACTCCC ACCGCCAAGG CCACCGGGGA GCCAAGGCC ACTAAACATA
 4601 TATGAAAAAA TGGCAACCGC TAATAAGGGG GCTATGACCG AAAATGCCA TGAAAACCGG CTACAGTCCTG ACCTAAAGG CAAACTGAT TCTGTCCTA
 ATACTTTT ACCGTTGCG ATTATCCCG CGATACTGGC TTGACCGCT ACCTTTGCGC GATGTCAGAC TGCGATTTC GTTGAACTA AGACAGCGAT
 4701 CTGATTACGG TGCTGCTATC GATGGTTCA TTGTTGACGT TCCGGCTT GCTATGGTA ATGTTGCTAC TGGTGAATTG GCTGGCTCTA ATTCCAAAT
 GACTAATGCC ACAGCGATAG CTACCAAAAGT AACCACTGCA AAGGGCGGA CGATTACCAT TACACGATG ACCACTAAA CGACCGAGAT TAAGGTTTA
 4801 GGCTCAAGTC GGTGACGGTG ATAATTCACT TTGTTGAAAT AATTCGTC AATATTACCC TTCTTTCGCT CAGTCGGTG AATGTCGCCCC TTATGTCCTT
 CCGAGITCAG CCACTGCCAC TATTAAGTGG AAATTACTTA TTAAGGCAG TTATAATGG AAGAAACGGA GTCAAGCCAC TTACAGGGG AATACAGAAA
 4901 GGCGCTGTTA AACCATATGA ATTTCCTATT GATGTTGACA AAATAAACTT ATTCCGTTG GTCTTTCGTT TTCTTATATA TTGTTGCCACC TTATGTA
 CCCGACCAT TTGGTATACT TAAAGATAA CTACACTGT TTGTTGAA TAAGGCACCA CAGAAACGCA AAGAAATAT ACAACGGTGG AAATACATAC

XbaI

Af III

5001 TATTTGAC GTTTGCTAAC ATACTGCGTA ATAAGGGACT TTAAGTAAT
 ATAAAAGCTG CAAACGATTG TATGACGCA TATTCTCAG AATTCATTA

FIG. 2B-3

10/522535

PCT/IB2003/003681

WO 2004/013276

Josef PRASSLER et al.
"Novel Tricistronic Vectors and Uses Therefor"
Attorney Docket No. 37629-0064US

6/25

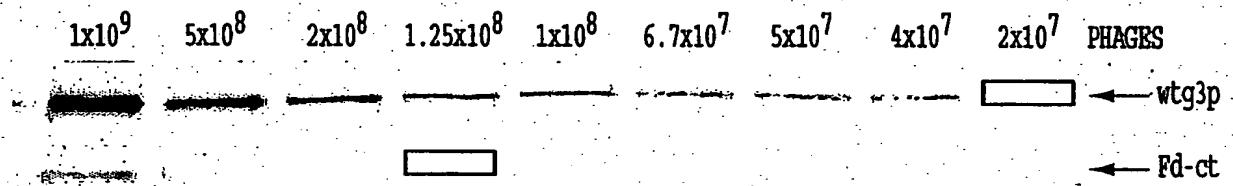


FIG. 3

10/522535

PCT/IB2003/003681

7/25

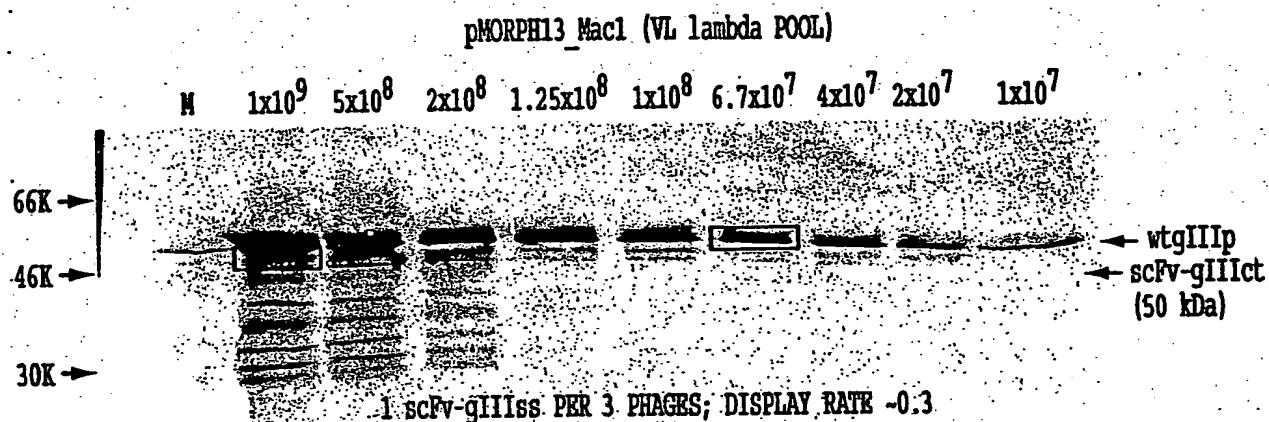


FIG. 4A

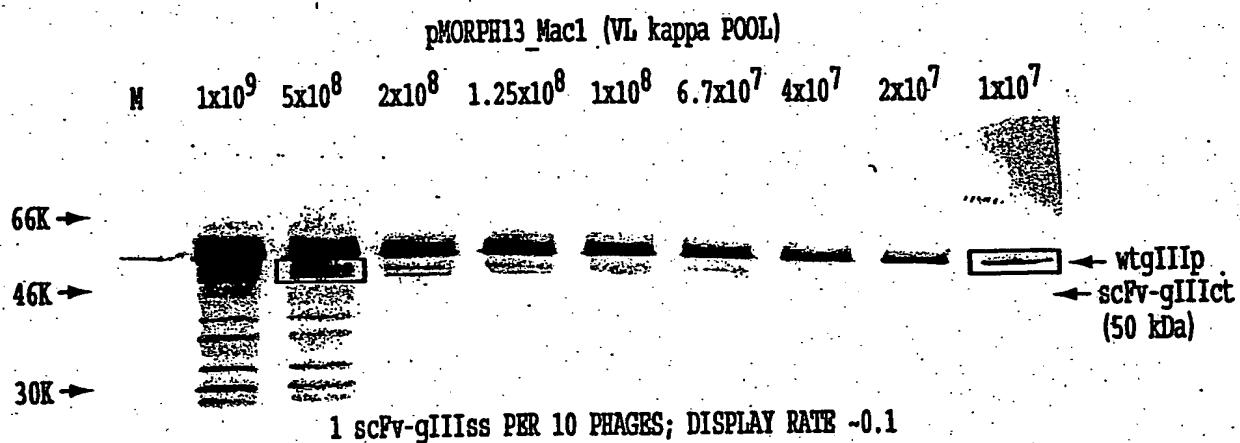


FIG. 4B

10/522535

WO 2004/013276

Josef PRASSLER et al.
 "Novel Tricistronic Vectors and Uses Therefor"
 Attorney Docket No. 37629-0064US

PCT/IB2003/003681

8/25

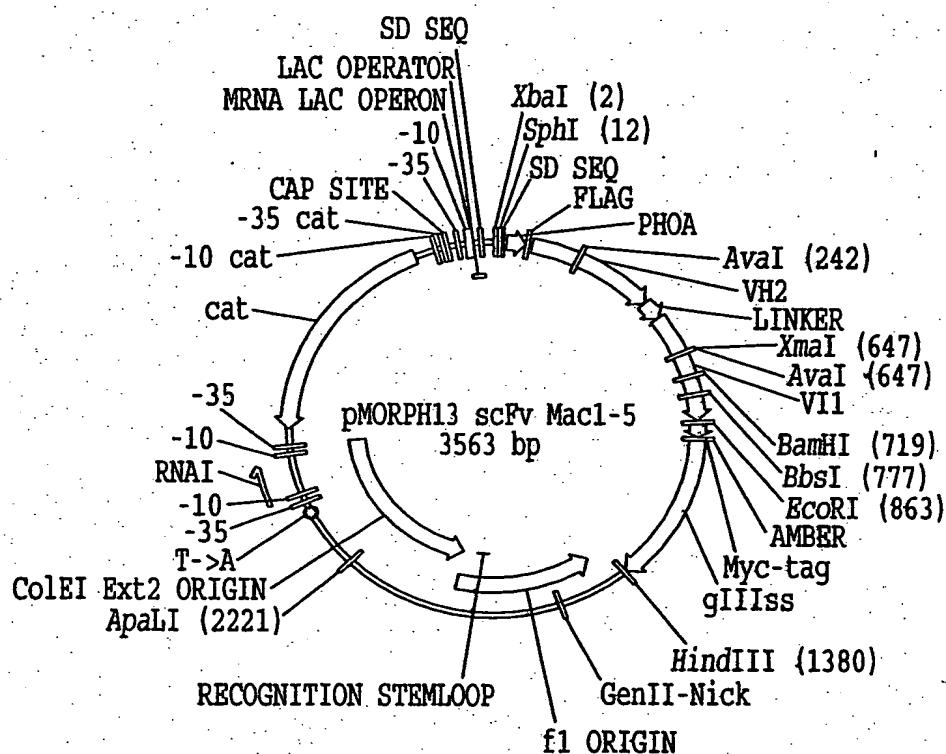


FIG. 4C

9/25

FIG. 4D

10/522535

WO 2004/013276

Josef PRASSLER et al.
"Novel Tricistronic Vectors and Uses Therefor"
Attorney Docket No. 37629-0064US

PCT/IB2003/003681

10/25

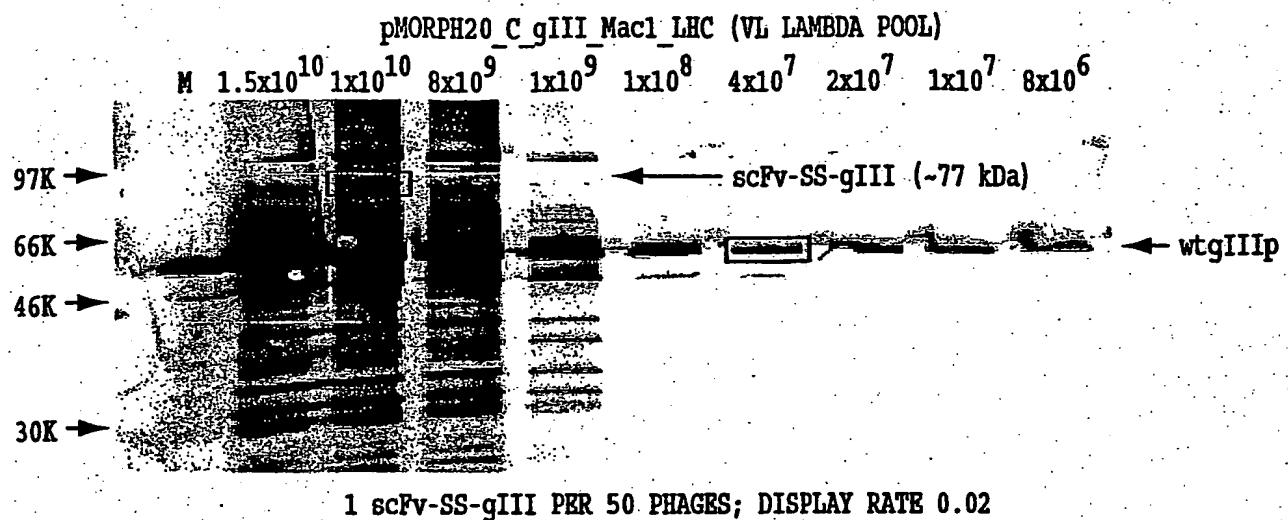


FIG. 5A

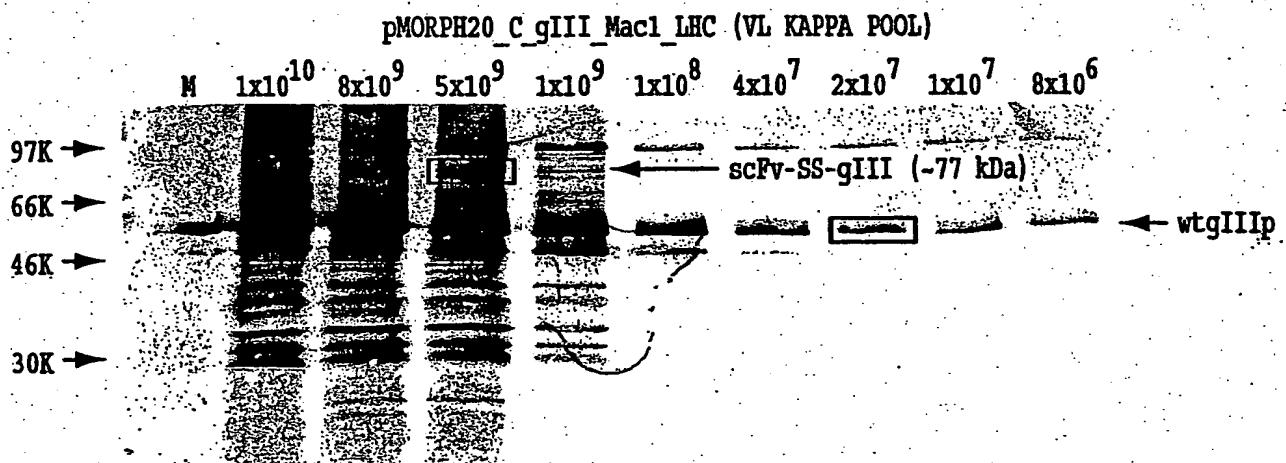


FIG. 5B

11/25

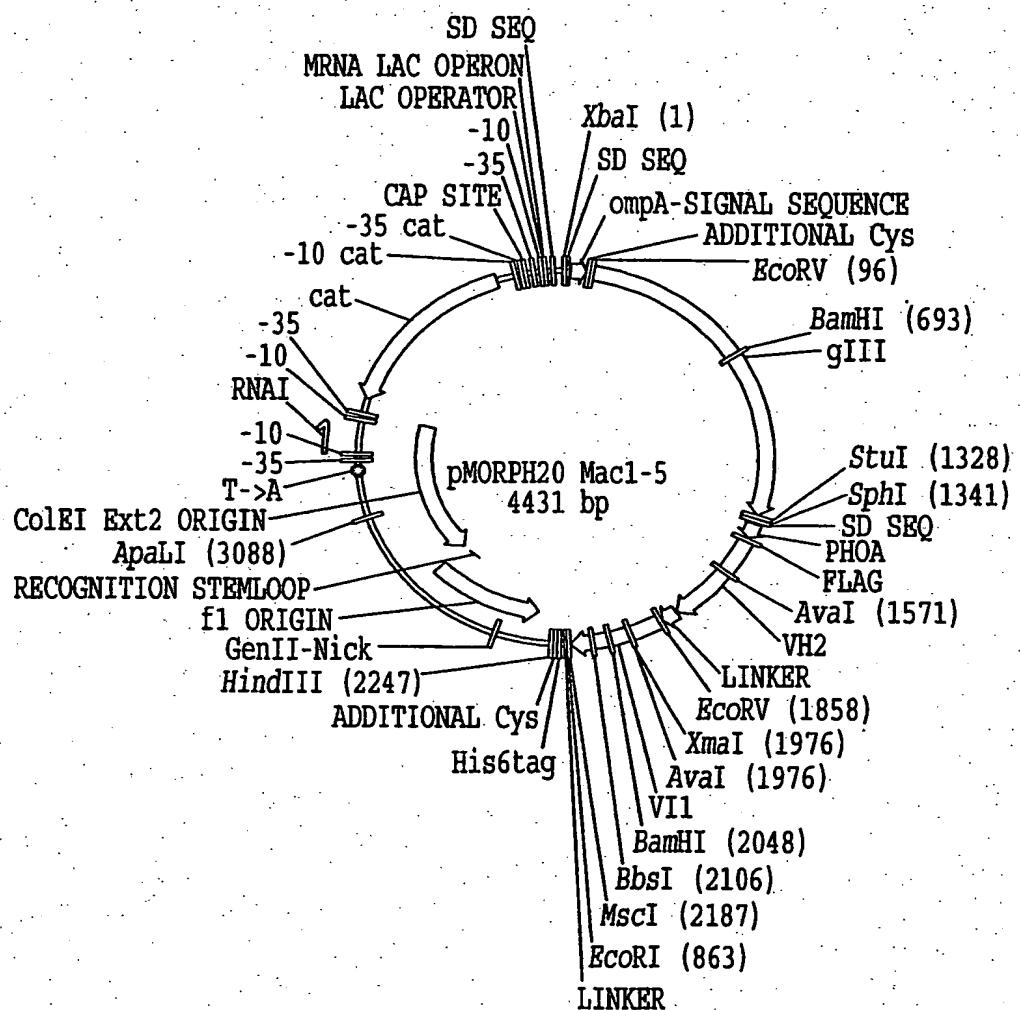
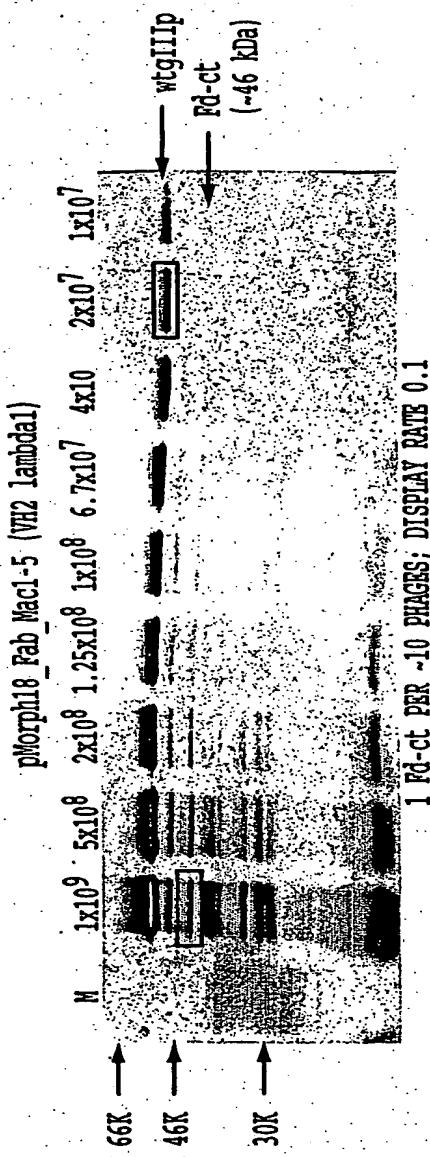
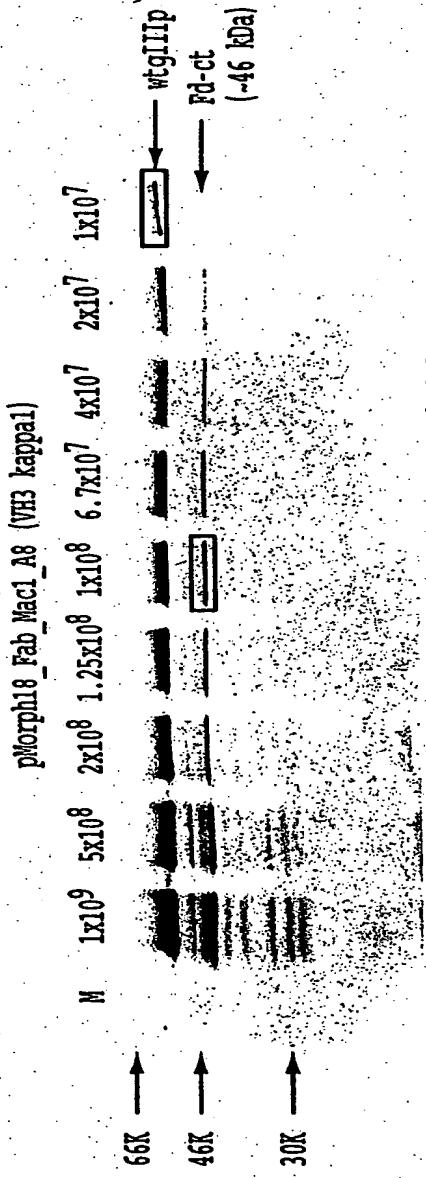


FIG. 5C

12/25

FIG. 5D

13/25

**FIG. 6A****FIG. 6B**

14/25

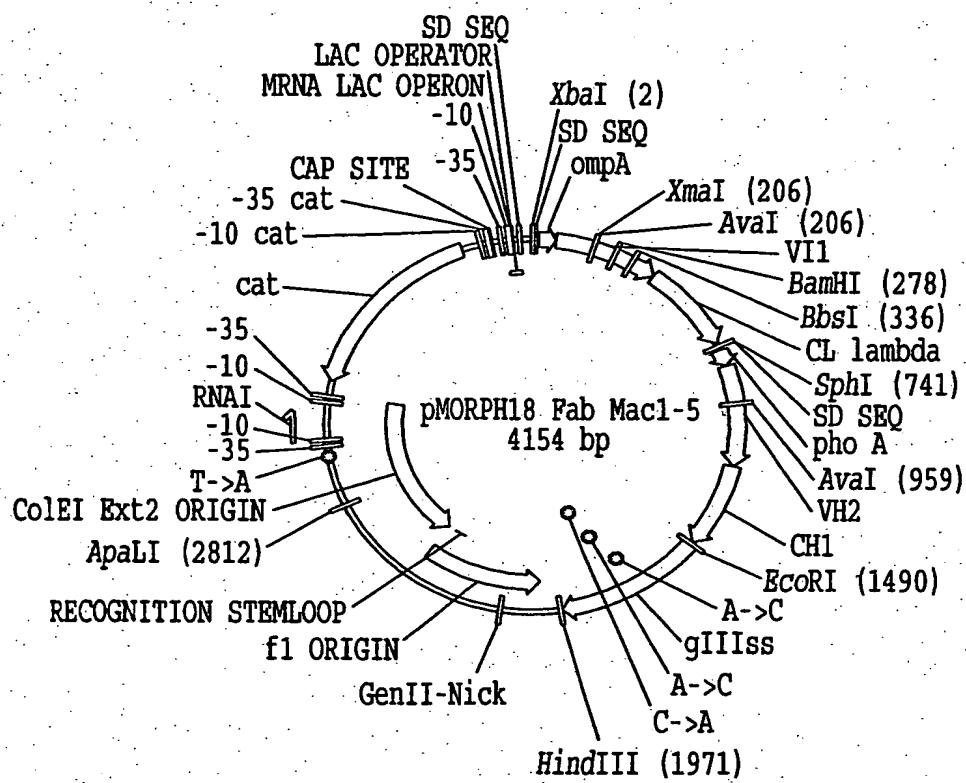


FIG. 6C

15/25

FIG. 6D

10/522535

WO 2004/013276

Josef PRASSLER et al.
"Novel Tricistronic Vectors and Uses Therefor"
Attorney Docket No. 37629-0064US

PCT/IB2003/003681

16/25

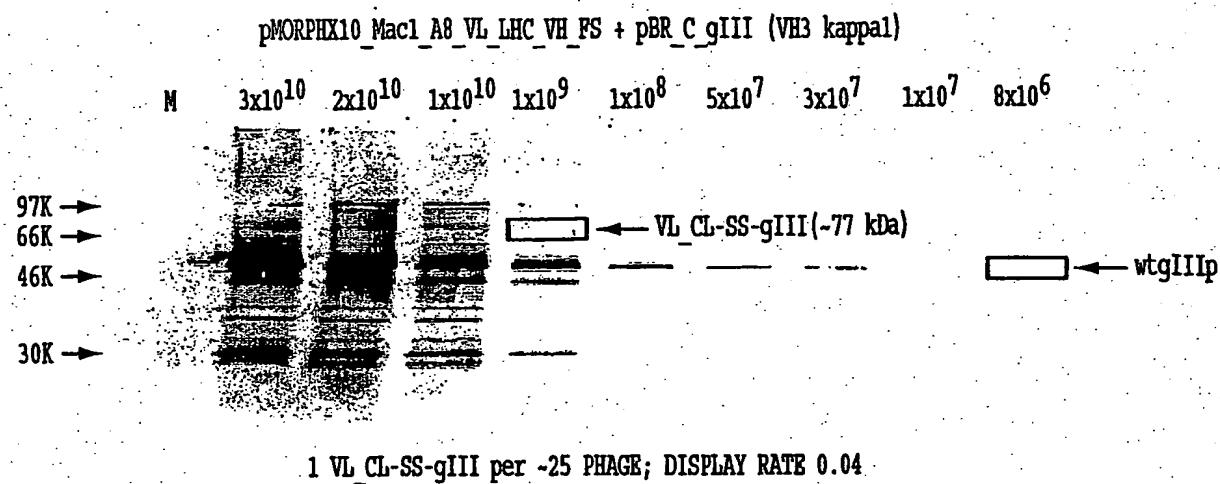


FIG. 7A

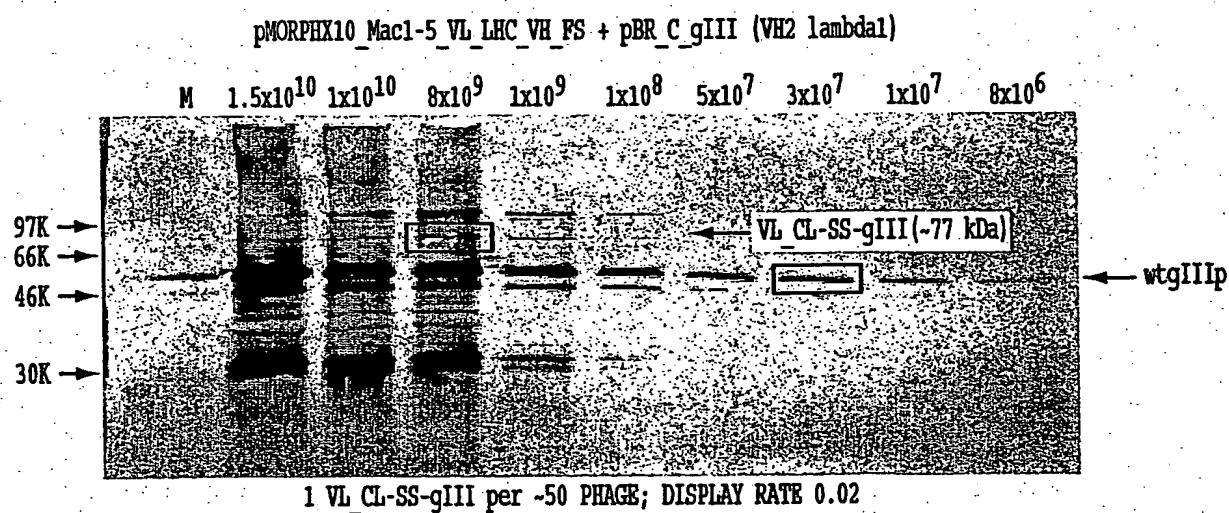


FIG. 7B

17/25

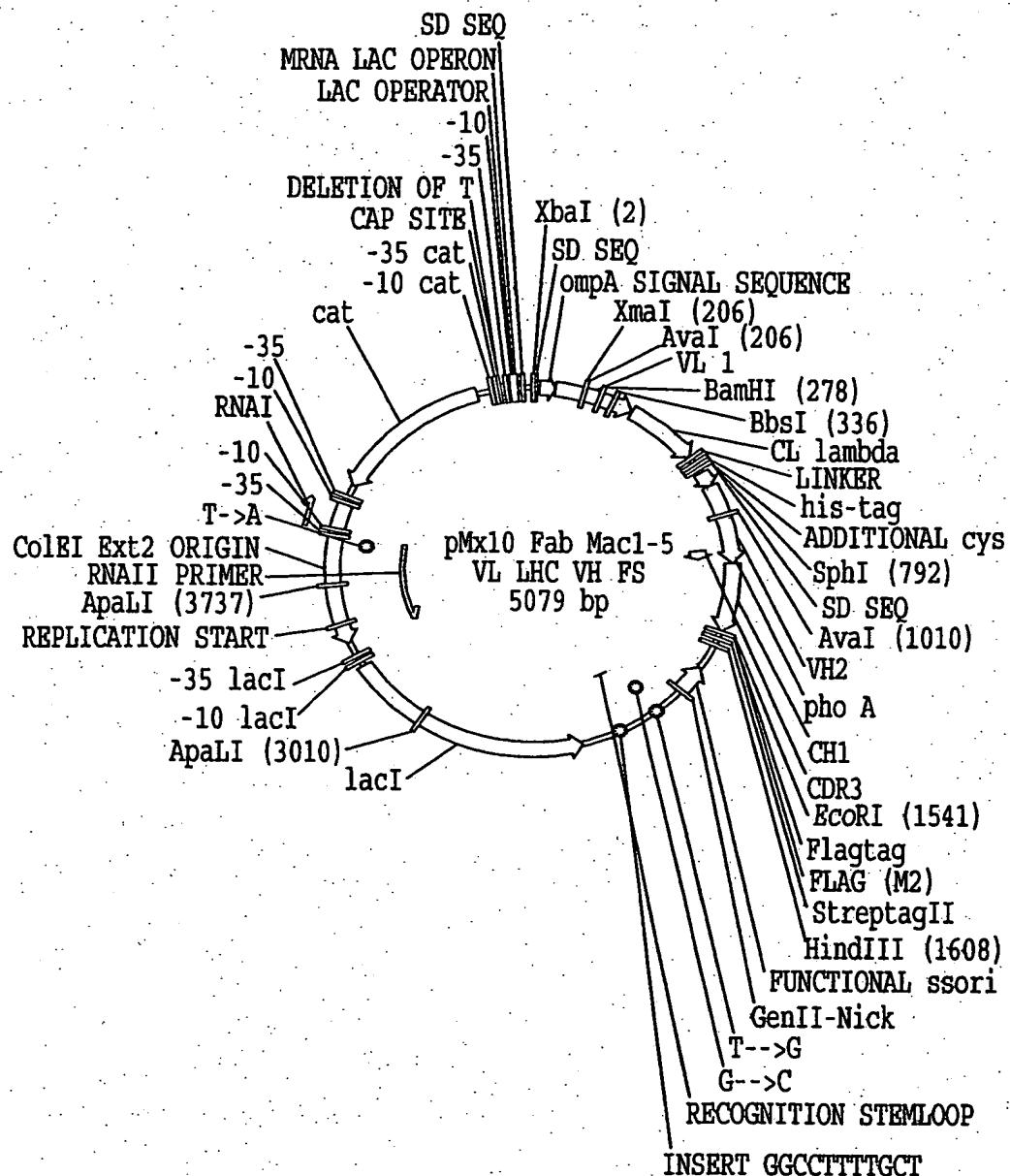


FIG. 7C

18/25

cagcagtgcggccggacggcgccqaaactgctgatttatgataacaaccagcgtccctcaggcgtgccggatcgtttagcggatc
 caaaaagcgccaccagcgcgagccgtcgattacgggctgcggcaaaagcgaagacgcaagcgattattattgcgcagagctatgacca
 gaatgctttgtgagggtttggcgccggcacaagttaccgttcttgccagccaaagccgaccgagtgtgacgctgtttcc
 gccgagcagcgaagaattcgaggcgaacaaagcaccctgggtgcctgattagcactttatccggagccgtgacagtggcc
 tggaggcagatagcggccgtcaaggcggagtgagacaccacccctccaaacaagaacaacaactacgcggccag
 cagctatctgagcctgacgcctgagcagtggagtcacccacagaagctacagctgcctgacgttgcacactgctgatgacgtggcc
 aaaaaccgttgcgcgactgaggccttcagggggagcggcgcggccaccatcatcaccatcactgctgataatatgca
 tgcgttaggagaaaataaaatgaaacaagcactattgcactggacttaccgttgatcttccctgttaccaaagcccagggt
 caattgaaagaaaagcgccggccgtggtaaaccgcacccctgaccctgacctgtacctttccggatttagcctgtccacg
 tctggcgttgcgtggctggattcgccagccctggaaagccctcgagtggctgtgattgttggatgtataatgt
 tatagcaccagcctgaaaacgcgtctgaccattagcaaagatacttcgaaaaatcagggtgtgtgactatgaccaacatggacc
 ggtggatacggccaccattattgcgcgtttgatctttttgttatt
 agctcagcgtcgaccaaggctcaagcgtttccgtggctccgagcagcacaaggcaccagcggggcacggctgcccctggct
 gcctggtaaagattttcccgaaaccagtaccgtgagctggaaacagcggggcgtgaccagcggcgtcataccctccgg
 gtgctgcaagcagcggccctgtatgcctgagcagcgttgcctgaccgtgcggagcagcacttaggcactcagactatattgca
 cgtgaaccataaaccgagcaacaccaagggtggataaaaatgg
 aggccgcgcgtggagcaccgcagggttggaaatgataacttgacctgtgacctgtgaaatgggggggggggggggggggggg
 tttgtctgcgtttaattaaagg
 tcgcgtttaatt
 gatagggttgagtgttgtccagtttggaaacaagacttgcactattaaqaacgtggactccaacgtctaaaggcggaaaaccgt
 atcaggggcgtggccactacgagaaccatcaccctaatcaagttttttgggtcgaggtgcgttacactaaatcggaaccc
 aaaggggagcccccgatttagcgttgcggggaaagccggcgaacgtggcgagaaggaaaggaaaggcggaaaaggagcg
 ggcgcctaggcgcgtggcaagtgtgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gctagactagtgtttaaaccggaccgg
 gggtagcctcactgcccgttccactcggaacccctgtcgccagctgcgtcatcgtgatcgcccaacgcgcggggagagg
 ttgcgttattggggccagggtgttttccactcggttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 ttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 cctcggtatcgtcgatccactaccgagatgtccgcaccaacgcgcggactcgtaatggcacgcattgcgttgcgttgcgt
 atctgatcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 atgcgccttcccggttccgttccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aacttaatggccagctaacgcgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gagaaaataatactgtgtatgggtgtcggttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 tagcatcctggatcagcggatgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 atggcccttcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gacgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 ataagagacaccggcatactctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 atgcccataccgcgaaagggttttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 cccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 acaggactataagataccagggttccctctggaaagctccctcgctgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 cgcctttccctcggttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gctgtgtgcacgaaccccccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aactacggctacactagaagaacagtattggatctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 cccggaaacaaaccaccgcgtgtgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 ccttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gggccaccaataactgccttaaaaaattacgcggccctgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 ggaagccatcacaacccgcgtatgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aaacggggcgaagaagtgtccatattggctacgtttaaatcaaaactggtaaactcaccaggatggctgagacgaaaa
 catattctaataaaaccctttagggaaataggcaggtttccctgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aaatcgtcggtatcactccagagcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 atcaccagctaccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aaacttgcctaaaatgttttacgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 aaatgccttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 ctgaaaatctcgataactaaaaatacccccgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gagttagctcactcattaggcacccccagggttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 acacaggaaacagctatgaccatgacattctgatgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gcaactggctgggttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgttgcgt
 gaccatctcggttagcggcagcagcaacatggcagcaactatgtgatgttgcgttgcgttgcgttgcgttgcgttgcgt

FIG. 7D

19/25

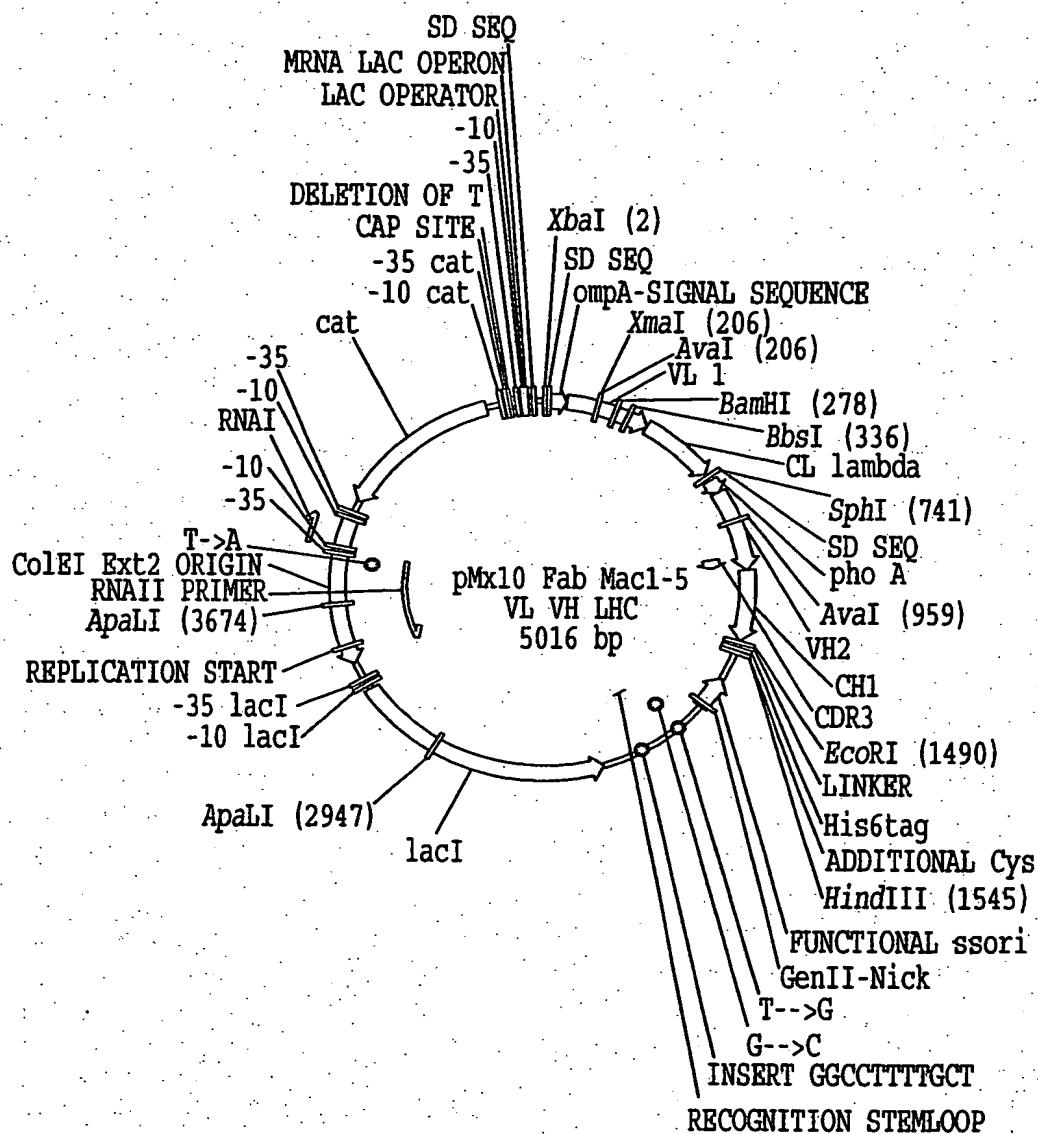


FIG. 7E

20/25

FIG. 7F

21/25

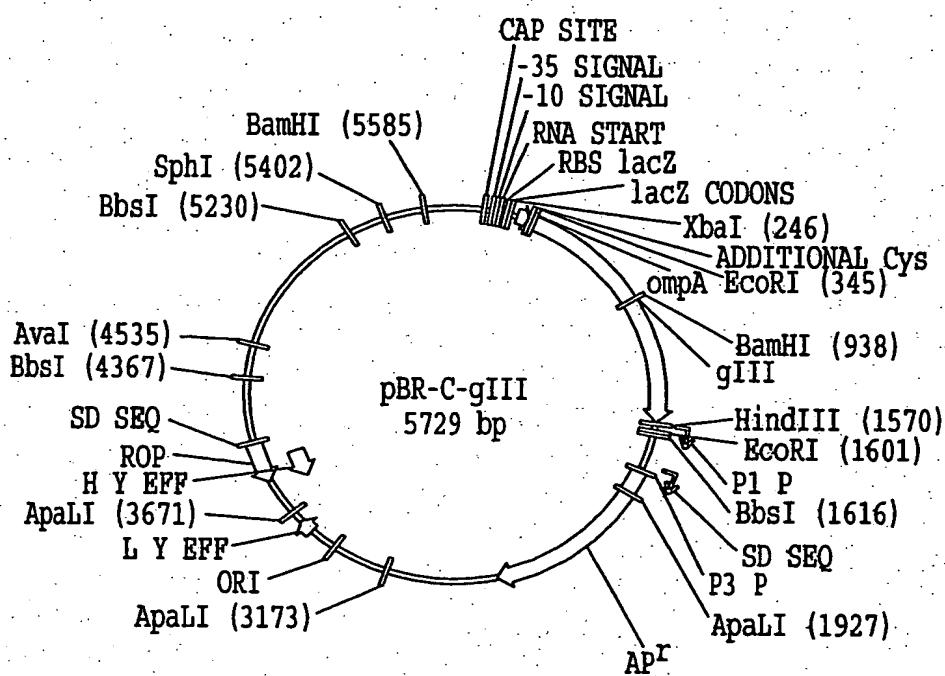


FIG. 7G

22/25

caagctgtaccgtctccggagctgcattgtcagaggtttaccgtcatcaccgaaacgcgcgaggcagtcggtaaagctc
 atcagcggtgtcgtaagcgattcacagatgtctgcctgttcatccgcgtccagctgttagtttcctccagaagcgtaatgtctggc
 ttctgataaagcgccatgttaagggcggttttccctgttgcactgtatgcctcgtaaggggatttcgttcatggggta
 atgataccgtgaaacgagagaggatgtcacgatcgggtactgtatgcatacatgcccggttactggAACGTTGTgagggt
 aacaactggcgatgtggatgcggcggaccagagaaaaatcaactcagggtcaatgccagcgcttcgttaatacagatgtagggt
 tccacagggtagccagcgcattctgcgtgcagatccggaacataatgtgcagggcgctgacttcgcgttccagacttacg
 aaacacgaaaccgaqaccattcatgttgttgcctaggcgcagcgtttgcagcagcagtcgttccacgttcgcgtatcg
 gtgattcattctgtctaaccgttaaggcaacccgcgcgcctagccgggtccgtcaacgcacaggagcactcatgcgcacccgtggc
 caggaccacgcgtcccgagatgcgcgcgtgcggctgctggagatgcggacgcgtggatatgttctgccaagggttggtt
 ggcattcacagtctccgcagaattgttgttgcattctggatgggtgatccgttagcggatgcggatccgcgttccatccatcg
 gtcgagggtggccggctccatgcaccgcacgcacgcggggaggcagacaaggatatagggcggccctacaatccatgcac
 ccgttccatgtgtccgcggcataatgcgggtccgtgcgttgcggatccgttgcggatccgcgttccatccatgcggccggaa
 ggcattcaagctgtccctgtatgggtgtcatctaccgcgtgcggatccgcgttgcggatccgcgttccatccatgcggccggaa
 gcgagaagaatcataatggggaaaggccatccagcgcgtgcggacgcgttgcggatccgcgttccatccatgcggcccatg
 ccggcgataatggcgttctcgccaaacgtttggccggggaccgttgcggatccgcgttccatccatgcggcccatg
 ataccgcacgcacaggccatcatgcgcgtccagcggaaacgggtccgtccgcgttgcggatccgcgttccatccatgcggcc
 tcctacgagttgcataaaagaagacagtcatgcggatccgcgttgcggatccgcgttgcggatccgcgttccatccatgcggcc
 ttgaaggctctcaaggcatcggtgcacgcgtcccttatgcactcgttgcatttaggaagcagcccagtagttaggttgcggcc
 agcaccgcgcgcgcacggatgtgcacgcgttgcggatccgcgttgcggatccgcgttccatccatgcggcc
 ccgaaacaagcgttgcacgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttccatccatgcggcc
 ctgtggccgggtgatgcggccacgtgcgtccgcgttagaggatccacaggacgggtgtggtgcggatccgcgttgcggcc
 agtggctcaagtgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggcc
 attgcataacgcataatgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 gcttcctgacaggaggccgtttgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 actttatgcgtccggctgtatgttgtgtggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 cttagataacgaggcaaaaatgaaaaagacagtatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ctactgcgtatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 cgacaaaactttagatgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tggtacggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tctgggggtggcggtactaaacgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 gcctggactgacaaaacccgcataatccgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ccgaaataggcggggcattaactgttataccggcactgttactcaaggactgcgttccattctggcttataacttcatgttgcggatccgc
 gtatcatccaaagccatgtatgcgttactgttataccggcactgttactcaaggactgcgttccattctggcttataacttcatgttgcggatccgc
 ttgtgaatataccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 gaggggtggcggtctggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ttgattatgaaaaatggcaaaacgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 caaactgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ttgtgattttgcgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tctttgcgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tccgtgggtgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 gcttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 taataatgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tgtatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ccttattccctttttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tgacgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 cactttaaagtgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 aatgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 catgagtgtataacactgcggccaaacttactgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 atcatgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 aatggcaacaacgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ataaagtgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 cggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 aacgaaatagacagatgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 tgatgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 ttccactgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 aaaaccaccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc
 gatccaatactgtccctgttgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgcgttgcggatccgc

FIG. 7H-1

SUBSTITUTE SHEET (RULE 26)

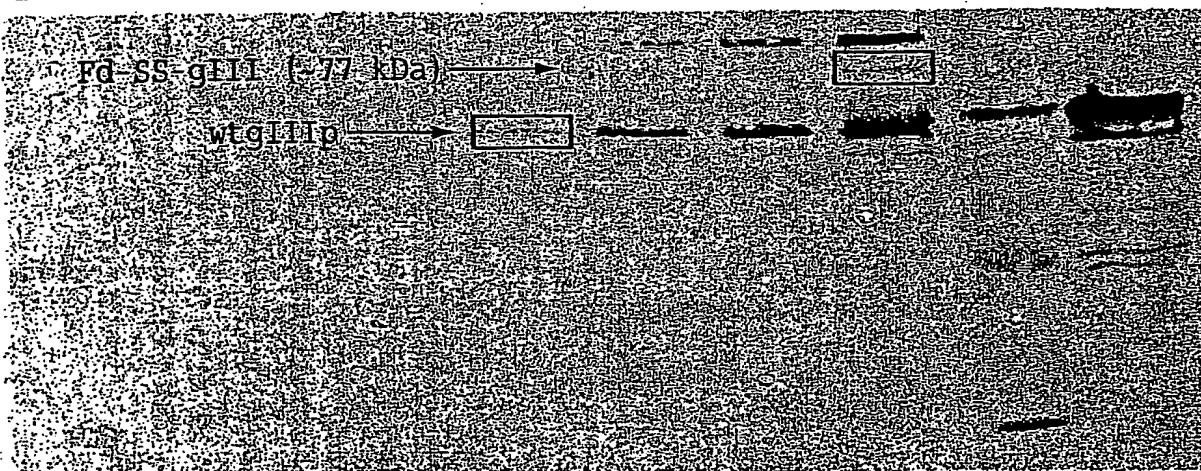
23/25

tcctgttaccagtggctgtgccagtggcgataagtcgtgttacccgggttgactcaagacgatagttaccggataaggcgacgc
ggtcgggctgaacgggggttcgtgcacacagccagttggagcgaacgacctacaccgaactcgagataccctacagcgtgagct
atgagaaaagcgccacgctcccgaaaggagaaaggcgacaggtatccggtaagcgccagggtcggaacaggagacgcacg
agggagcttcaggggaaacgcctgttatctttatagtcgtcggttgcgcacctctgacttgagcgtcgatttgtatgctc
gtcaggggggcggagctatggaaaaacgcccagcaacgcggcttttaeaggttctggcctttgtcgccctttgtcacatgttc
tttcctgcgttatccccgtattctgtggataaccgtattaccgccttgagttagtgcataccgcctcgccgcagccqaacgaccgacg
gcagcgagtcagtgagcgagaagcggaagagcgccgtatgcgtatttctccatgcacatctgtcggtattcacaccgatcat
ggtgactctcgtacaatctgctgtatgcgcatagttaaagccagtatacactccgcatacgactggcatggctcg
ccccgacacccgccaacacccgctgacgcgcctgtctgcctccggcatccgcttacaga

FIG. 7H-2

24/25

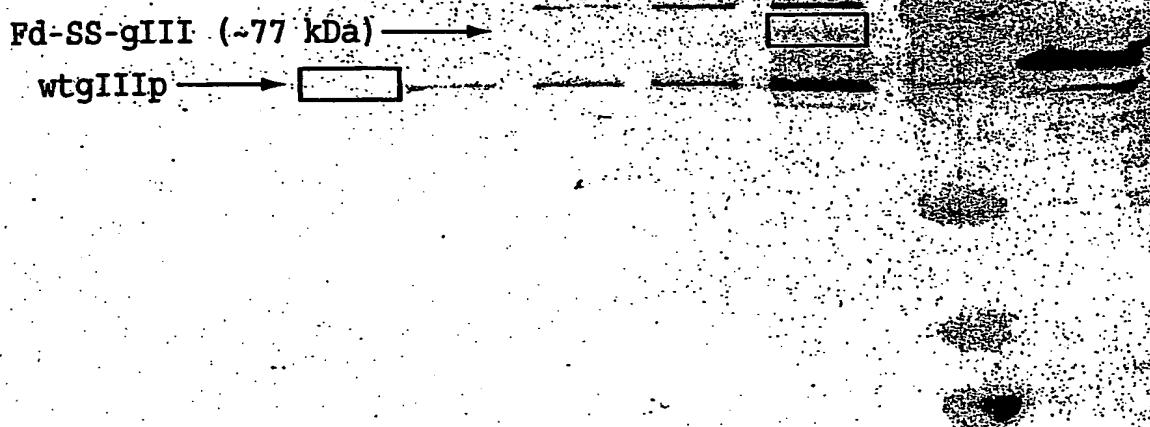
pMORPH23_VH3_Vk+1 (pool)

 1×10^8 2×10^8 5×10^8 1×10^9 3×10^9 1×10^{10} 2×10^{10} 5×10^{10} M +DTT; 5×10^9


1 Fd-SS-gIII PER ~3 PHAGE; DISPLAY RATE 0.3

FIG. 8A

pMORPH23_VH2_Vk+1 (pool)

 1×10^7 2×10^7 5×10^7 1×10^8 3×10^8 1×10^9 2×10^9 5×10^9 M +DTT; 5×10^9


1 Fd-SS-gIII PER ~10 PHAGE; DISPLAY RATE 0.1

FIG. 8B

25/25

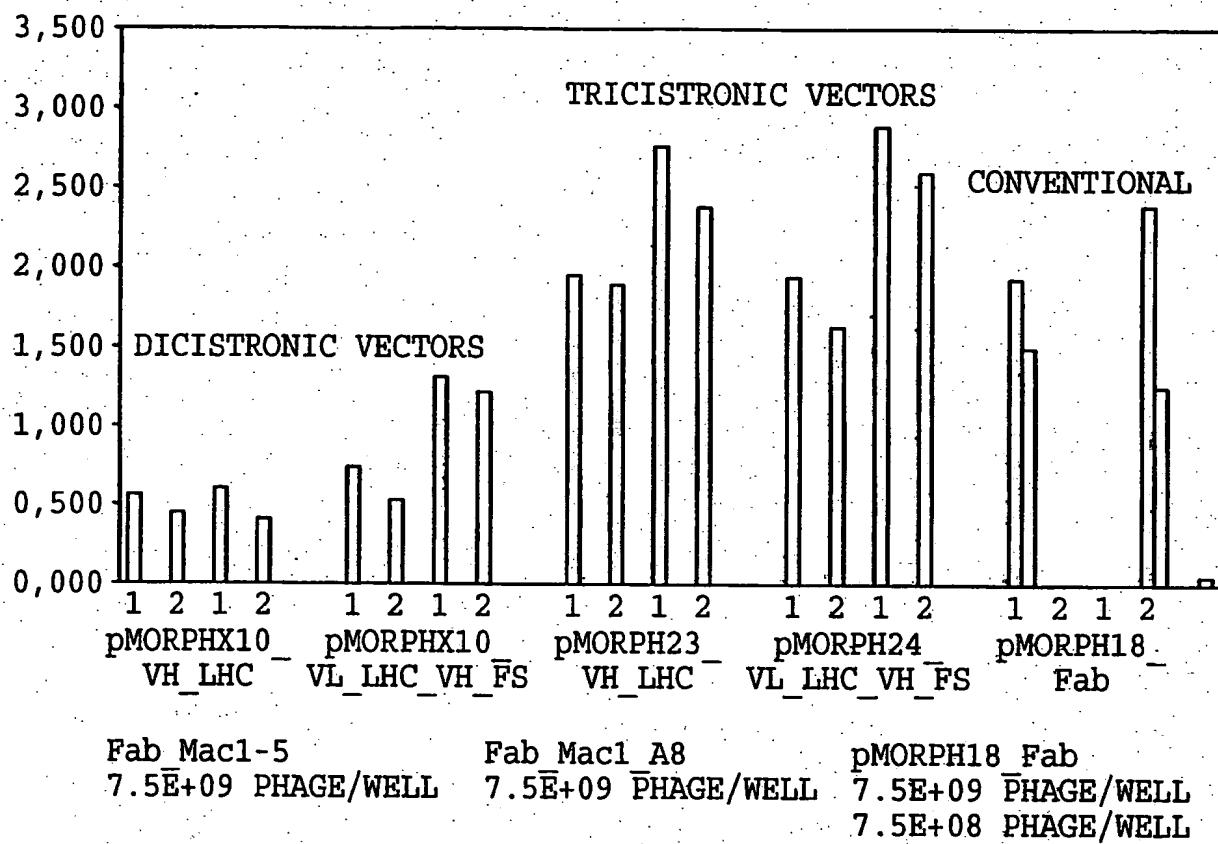


FIG. 9